Serial No. 09/158,982 Response to Office Action of Feb. 5, 2003

Amendments to the Claims

Please cancel Claim 21 and amend Claims 3, 6, 7, 11-14 and 23 as follows:

- 3. (Currently amended) A method in a computer system for analyzing an and displaying data on gene expression in a molecular topography, comprising:
- (a) generating a gene expression profile of a plurality of gene-expression indicating polynucleotides including for each of the polynucleotides:
 - (1) a first value for a first polynucleotide characteristic, comprising a sequence identifier;
 - (ii) a second value for a second polynucleotide-characteristic different from said first characteristic; comprising a measure of size of the polynucleotide; and
 - (iii) a third value that is a measure of the quantity of the polynucleotide;
- (b) calculating for each polynucleotide from the first and second values a position, and from the third value a peak at that position in a multi-dimensional display space; and
- displaying the peak for each polynucleotide at the calculated position for the polynucleotide in the display, the resulting display representing thereby a molecular topography of gene expression;
- wherein: (i) the first-characteristic is a sequence identifier, and (ii) the second characteristic is a measure of size.
- 6 (Currently amended) A method in a computer system for analyzing an and displaying data on gene expression in a molecular topography, comprising:
- (a) generating a gene expression profile of a plurality of gene-expression indicating polynucleotides, wherein the polynucleotides are 3'-end fragments of restriction enzyme cleaved cDNAs or fragments thereof, the gene expression profile including for each of the polynucleotides:
 - (i) a first value for a first polynucleotide characteristic, comprising a sequence identifier;
 - (ii) a second value for a second polynucleotide-characteristic different from said first characteristic; comprising a measure of size of the polynucleotide; and
 - (iii) a third value that is a measure of the quantity of the polynucleotide;

ATLLIBOT 1439932 F (2002 K-4)

- (b) calculating for each polynucleotide from the first and second values a position, and from the third value a peak at that position in a multi-dimensional display space; and
- (c) displaying the peak for each polynucleotide at the calculated position for the polynucleotide in the display, the resulting display representing thereby a molecular topography of gene expression;

wherein the polynucleotides are cDNAs or fragments thereof, and
wherein the polynucleotides are 3' end fragments of restriction enzyme cleaved cDNAs.

- 7. (Currently amended) The method of claim 6, wherein the first characteristic is a sequence identifier, and the second characteristic is a measure of size of the fragment.
- 8. (Original) The method of claim 7, wherein the sequence identifier is the combination of an anchor sequence of a cDNA synthesis primer and a restriction enzyme cleavage reaction specificity.
- 9. (Original) The method of claim 8, wherein values of the first and second characteristics are arranged, respectively, along first and second ordinates, the first and second ordinates defining a plane, the values of the first and second characteristics for each polynucleotide by their location on the first and second ordinates define a position in the plane, and the peak for each polynucleotide is displayed at the position of the polynucleotide on the plane.
- 10. (Original) The method of claim 9, wherein the 3' end fragments are generated by READS.
- 11. (Currently amended) A method in a computer system for displaying differences in gene expression, comprising:
- (1) for each of two gene expression profiles a first gene expression profile and a second gene expression profile to be compared, generating a molecular topography according

ATULI801 1439932 1 [2002 K14]

to a method in a computer system for analyzing an and displaying data on gene expression in a molecular topography, comprising:

- (a) generating a gene expression profile of a plurality of gene-expression indicating polynucleotides including for each of the polynucleotides:
 - (i) a first value for a first polynucleotide characteristic,
 - (ii) a second value for a second polynucleotide-characteristic different from said first characteristic, and
 - (iii) a third value that is a measure of the quantity of the polynucleotide;
- (b) calculating for each polynucleotide from the first and second values a position in a two co-ordinate display space, and from the third value a peak at that position in a multi-dimensional display-space; and
- (c) displaying the peak for each polynucleotide at the calculated position for the polynucleotide in the display, the resulting display representing thereby a molecular topography of gene expression,
- (2) for each position in the two co-ordinate <u>display</u> space subtracting the measure of quantity in the first <u>gene expression</u> profile from the measure of quantity in the second <u>gene expression</u> profile;
 - (3) for each position, generating a peak for display from the difference; and
- (4) displaying the respective difference peak at each of the positions in the two coordinate <u>display</u> space.
- 12. (Currently amended) A method in a computer system for displaying differences in gene expression, comprising:
- (1) for each of two gene expression profiles a first gene expression profile and a second gene expression profile to be compared, generating a molecular topography according to a method, comprising:
 - (a) generating a gene expression profile of a plurality of gene-expression indicating polynucleotides including for each of the polynucleotides:
 - (i) a first value for a first polynucleotide characteristic,

ATLLIB01 1439932 1 [2002 K14]

- (ii) a second value for a second polynucleotide-characteristic different from said first characteristic; and
- (iii) a third value that is a measure of the quantity of the polynucleotide;
- (b) calculating for each polynucleotide from the first and second values a position in a two-co-ordinate display space, and from the third value calculating a peak at that position in a multi-dimensional display-space; and
- (c) displaying the peak for each polynucleotide at the calculated position for the polynucleotide in the display, the resulting display representing thereby a molecular topography of gene expression;
- (2) for each position in the two co-ordinate <u>display</u> space subtracting the measure of quantity in the first <u>gene expression</u> profile from the measure of quantity in the second <u>gene expression</u> profile;
 - (3) for each position, generating a peak for display from the difference; and
- (4) displaying the respective difference peak at each of the positions in the two coordinate display space,

wherein differences are displayed only if they meet or exceed a threshold value.

- 13. (Currently amended) A method in a computer system for comparing gene expression, comprising:
- (1) for each of two gene profiles a first gene expression profile and a second gene expression profile to be compared, generating a molecular topography according to claim ± 11.
- (2) for each position in the two co-ordinate <u>display</u> space subtracting a first measure of quantity in the first profile from a second measure of quantity in the second profile to generate respective difference values;
- (3) calculating from the respective difference values a measure of difference of the two profiles.
- 14. (Currently amended) The method of claim 13, wherein the first profile is a standard reference profile and the second profile is a the profile of a test sample.

ATULIB01 1439932 1 [2002 K14]

- 15. (Original) The method of claim 14, wherein comparison of the test sample profile with the standard reference profile is diagnostic of a condition or disease.
- 16. (Original) The method of claim 15, wherein the first profile is a profile of a reference sample and the second profile is a profile of a test sample.
- 17. (Original) The method of claim 16, wherein comparison of the profile of the test sample with the profile of the reference sample is diagnostic of a condition or disease.
- 18. (Original) The method of claim 13 wherein the difference value is set to zero when the difference between the first measure and the second measure is less than two-fold.
- 19. (Original) The method of claim 13 wherein the first profile is identical to the second profile when all the respective first and second measures are identical.
- 20. (Original) The method of claim 13 wherein the first profile is similar to the second profile when the average difference between the respective first and second measures is less than a predefined amount.
 - 21. (Cancelled)
- 23. (Currently amended) A method in a computer system for displaying a series of in gene expression profiles in a molecular movie, comprising:
- (1) generating a plurality of molecular topographies according to a method, comprising:
 - (a) generating a gene expression profile of a plurality of gene-expression indicating polynucleotides including for each of the polynucleotides:
 - (i) a first value for a first polynucleotide characteristic, comprising a sequence identifier;
 - (ii) a second value for a second polynucleotide-characteristic different from said first characteristic, comprising a measure of size of the polynucleotide; and

ATLLIB01 1439932 | [2002 K14]

Scrial No. 09/158,982 Response to Office Action of Feb. 5, 2003

- (iii) a third value that is a measure of the quantity of the polynucleotide;
- (b) calculating for each polynucleotide from the first and second values a position, and from the third value a peak at that position in a multi-dimensional display space; and
- (c) displaying the peak for each polynucleotide at the calculated position for the polynucleotide in the display, the resulting display representing thereby a molecular topography of gene expression; and
- (2) displaying the molecular topographies in succession; wherein the molecular topographies are delta plots.